SEQUENCE LISTING

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<110> Tsuchiya, Masayuki
     Saito, Mikiyoshi
     Ohtomo, Toshihiko
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Gly Ala Ala Leu Ala Pro Arg Arg Cys Pro Ala Gln Glu Val Ala Arg
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                              25
                                                                  144
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Gly Val Leu Thr Ser Leu Pro Gly Asp Ser Val Thr Leu Thr Cys Pro
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								gga Gly		240
								tat Tyr		288
								gtg Val 110		336
	 	_		_		_	_	ccc Pro	_	384
								ctg Leu		432
								gcc Ala		480
								ttc Phe		528
								gtg Val 190		576
								caa Gln		624
								atc Ile		672
								tgg Trp		720
								gag Glu		768
								gtc Val 270		816

ctc cag cat cac tgt gtc atc cac gac gcc tgg agc ggc ctg agg cac 864 Leu Gln His His Cys Val Ile His Asp Ala Trp Ser Gly Leu Arg His 275 912 gtg gtg cag ctt cgt gcc cag gag gag ttc ggg caa ggc gag tgg agc Val Val Gln Leu Arg Ala Gln Glu Glu Phe Gly Gln Gly Glu Trp Ser 290 295 300 gag tgg agc ccg gag gcc atg ggc acg cct tgg aca gaa tcc agg agt 960 Glu Trp Ser Pro Glu Ala Met Gly Thr Pro Trp Thr Glu Ser Arg Ser 315 310 cct cca gct gag aac gag gtg tcc acc ccc atg cag gca ctt act act 1008 Pro Pro Ala Glu Asn Glu Val Ser Thr Pro Met Gln Ala Leu Thr Thr 325 330 aat aaa gac gat gat aat att ctc ttc 1035 Asn Lys Asp Asp Asn Ile Leu Phe 340 345 <210> 3 <211> 40 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: "IL6R1", an artificially synthesized primer sequence <400> 3 ttcgaattcc caccatgctg gccgtcggct gcgcgctgct 40 <210> 4 <211> 36 <212> DNA <213> Artificial Sequence <223> Description of Artificial Sequence: "IL6R2", an artificially synthesized primer sequence 36 ttcgaattcg aagagaatat tatcatcgtc tttatt <210> 5 <211> 768 <212> DNA <213> Artificial Sequence <220> <221> CDS <222> (1)..(768) <220> <223> Description of Artificial Sequence: a designed single chain Fv gene sequence

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	ctg Leu								96	
	gcc Ala								144	
	gga Gly 50			_					192	
	tcc Ser								240	
	aga Arg								288	
	aga Arg								336	
	ctc Leu								384	
	ggt Gly 130								432	
	agc Ser								480	
	gac Asp								528	
	cca Pro								576	
	agc Ser								624	
	agc Ser 210								672	

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Gly Asn Thr Leu Pro Tyr Thr Phe Gly Gln Gly Thr Lys Val Glu Ile
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aaa cga act gtg gct gca cca tct gtc ttc atc ttc ccg cca tct gat
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•																	
											8						
	aac	tca	aac	ttc	act	qcc	att	tat	ata	ctq	aaq	qaq	qcq	tgt	ctg	cag	256
							Ile										
					45					50					55		
						~~~	200	<b>+</b>	250	~+~	+ ~~	224	200	226	ast	aat	304
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		-1-	-1-	60				-1-	65			-1-	<del></del> -	70			
							gtc										352
	ALA	var	75	Arg	GIU	GIN	Val	80	vaı	TTE	ASI	Arg	85	Thr	ser	ser	
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	gtc	acg	ttc	aca	gac	gtg	gtc	ctc	ccg	agc	gtg	cag	ctc	acc	tgc	aac	400
-	Val		Phe	Thr	Asp	Val	Val	Leu	Pro	Ser	Val		Leu	Thr	Суз	Asn	
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	105					110					115					120	
	ctt	tca	aac	+++	cct	cca	gat	222	cct	aca	aat	tta	act	tac	att	ata	496
							Asp					_		_			150
			_		125		_	-		130				<u> </u>	135		
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		Pro	Thr	Tyr	Tyr		Asn	Ile	Glu	Val	_	Val	Glu	Ala	Glu		
	185					190					195					200	
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	222	ata.	222		200	000	cca	+-+	224	++-	+ 00	~+~	200	220	tas	<b>~</b> ~~	784
							Pro									_	
	-1-		-4-	220				-1-	225					230			
							aag										832
	GLU	⊥eu	235	Ser	TTE	neu	Lys	240	ser.	тър	val	SEL	245	GTĀ	neu	GTÀ	
				_		_	tct	_							_	_	880
	Gly		Leu	Asp	Leu	Lys	Ser	Asp	Ile	Gln	Tyr	_	Thr	Lys	Asp	Ala	
		250					255					260					

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													gtg Val			9	76
				_		_							gac Asp 310			10	24
													aga Arg			10	72
													tat Tyr			11	.20
_					_	_	_				_	_	aat Asn			11	.68
													gtc Val			12	16
													acc Thr 390			12	64
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													tgg Trp			14	80.
									_				gag Glu			14	56
	_				~		_	_	_	_		-	cag Gln 470	_	_	15	04
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tgc	tat	caa	atc	aca	gta	act	ccc	gta	ttc	gcc	acg	<b>9</b> 99	ccc	gga	ggc	16	00

Cys	Tyr 490	Gln	Ile	Thr	Val	Thr 495	Pro	Val	Phe	Ala	Thr 500	Gly	Pro	Gly	Gly	
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					aag Lys											1696
	_	_			gtg Val	_	_	_								1744
					acc Thr											1792
					gag Glu											1840
					atg Met 590											1888
					ttt Phe											1936
			-		cct Pro		_		-			_			_	1984
					tgc Cys											2032
					cct Pro											2080
					cca Pro 670											2128
					ttc Phe											2176
					tgt Cys											2224
					agt Ser											2272

715 720 725

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						tcc act gtg Ser Thr Val 760	
						gtg ttc tca Val Phe Ser 775	
				Asp Ser		cgg cca gaa Arg Pro Glu 790	
	Leu Val	_				atc ttg ccc Ile Leu Pro	
			_		_	gcc tgt cca Ala Cys Pro	
						ggc aat gag Gly Asn Glu 840	
	-				_	cac att tct His Ile Ser 855	
				Leu Phe		ggc tct aca Gly Ser Thr 870	
	Leu Gly					aga ttt gaa Arg Phe Glu	
			Ile Asp			aaa agt tac Lys Ser Tyr	
ttg cca cag Leu Pro Gln 905						tgaaggactg	2849
gctcctgaac	ttcagcag	ga actgc	aaaat aa	agctaaag	acgagtg	gct tcagatga	ga 2909
aacagtcctc	actccctg	aa gatag	gcatt go	ctctaagg	acaaagt	cac acctgggc	cg 2969
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                                                                   97
get aca ggt gtc gac tec cag gtc caa etg cag gag age ggt eca ggt
Ala Thr Gly Val Asp Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly
     15
                         20
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ctt Leu 30	gtg Val	aga Arg	cct Pro	agc Ser	cag Gln 35	acc Thr	ctg Leu	agc Ser	ctg Leu	acc Thr 40	tgc Cys	acc Thr	gtg Val	tct Ser	ggc Gly 45	145
							gcc Ala									193
							gga Gly									241
							tcc Ser 85									289
~	_		_		_	_	aga Arg		_				_		-	337
							aga Arg									385
							ctc Leu									433
							ggt Gly									481
							agc Ser 165									529
							gac Asp									577
							cca Pro									625
							agc Ser									673
							agc Ser									721
			_		_		aac Asn 245	-								769
ggg	acc	aag	gtg	gaa	atc	aaa	tct	aga	ggt	ggt	ggt	ggt	tcg	ggt	ggt	817

Gly	Thr 255	Lys	Val	Glu	Ile	Lys 260	Ser	Arg	Gly	Gly	Gly 265	Gly	Ser	Gly	Gly	
														ctg Leu		865
														ctg Leu 300		913
														agc Ser		961
														att Ile		1009
														gtg Val		1057
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														cta Leu 380		1153
														aca Thr		1201
						_					_			ggc Gly		1249
														agc Ser		1297
														agc Ser		1345
														ctg Leu 460		1393
					_	_						_	_	ttc Phe	_	1441
														ctc Leu		1489

485

480

490

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					tgg Trp											192
					tac Tyr 70											240
					aga Arg											288
					ctc Leu											336
					tcc Ser											384
					gtc Val											432
					ggt Gly 150											480
					gcc Ala											528
Arg	Ala	Ser	Gln 180	Asp	atc Ile	Ser	Ser	Tyr 185	Leu	Asn	Trp	Tyr	Gln 190	Gln	Lys	576
Pro	Gly	Lys 195	Ala	Pro	aag Lys	Leu	Leu 200	Ile	Tyr	Tyr	Thr	Ser 205	Arg	Leu	His	624
Ser	Gly 210	Val	Pro	Ser	aga Arg	Phe 215	Ser	Gly	Ser	Gly	Ser 220	Gly	Thr	Asp	Phe	672
					agc Ser 230											720
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	gaa Glu															780

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                                                                   96
Gln Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe
                                                                   144
tat ccc aga gag gcc aaa gta cag tgg aag gtg gat aac gcc ctc caa
Tyr Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln
                             40
teg ggt aac tee cag gag agt gte aca gag cag gae age aag gae age
                                                                   192
Ser Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser
acc tac agc etc agc agc acc etg acg etg agc aaa gca gac tac gag
                                                                   240
Thr Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu
                                         75
                                                                   288
aaa cac aaa gtc tac gcc tgc gaa gtc acc cat cag ggc ctg agc tcg
Lys His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser
                 85
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ccc gtc aca aag agc ttc aac agg gga gag tct
Pro Val Thr Lys Ser Phe Asn Arg Gly Glu Ser
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Val Ala Leu His Arg Pro Asp Val Tyr Leu Leu Pro Pro Ala Arg Glu
cag ctg aac ctg cgg gag tcg gcc acc atc acg tgc ctg gtg acg ggc
                                                                    96
Gln Leu Asn Leu Arg Glu Ser Ala Thr Ile Thr Cys Leu Val Thr Gly
             20
                                  25
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ttc tct ccc gcg Phe Ser Pro Ala 35	gac gtc ttc Asp Val Phe	gtg cag tgg Val Gln Trp 40	atg cag agg ggg c Met Gln Arg Gly G 45	ag ccc 144 Eln Pro
			cca atg cct gag of Pro Met Pro Glu F	
			ctg acc gtg tcc g Leu Thr Val Ser G 75	
			gtg gcc cat gag g Val Ala His Glu <i>F</i>	
	Thr Glu Arg		aag tee ace gag g Lys Ser Thr Glu G 110	
gtg agc gcc gac Val Ser Ala Asp 115				363
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			gtg tct ggc tac t Val Ser Gly Tyr S 45	
			cag cca cct gga c Gln Pro Pro Gly F 60	

				gga Gly												240
				tcc Ser 85	-			_		_	-		-	_		288
				aga Arg		-	-	-		_	_	_				336
				aga Arg												384
				ctc Leu												432
				ggt Gly												480
				agc Ser 165												528
-	_	_	_	gac Asp		_	~		_				_	_	_	576
				cca Pro												624
				agc Ser												672
				agc Ser												720
				aat Asn 245												768
				cga Arg												816
				cag Gln												864
ctg	aat	aac	ttc	tat	ccc	aga	gag	gcc	aaa	gta	cag	tgg	aag	gtg	gat	912

Leu Asn Asn												
290	Phe Tyr	Pro Ar 29		Ala	Lys	Val	Gln 300	Trp	Lys	Val	Asp	
aac gcc ctc												960
Asn Ala Leu	Gln Ser	-	n Ser	Gln	Glu		Val	Thr	Glu	Gln	_	
305		310				315					320	
agc aag gac	agc acc	tac ag	c ctc	agc	agc	acc	ctg	acg	ctg	agc	aaa	1008
Ser Lys Asp		Tyr Se	r Leu	Ser		Thr	Leu	Thr	Leu		Lys	
	325				330					335		
gca gac tac	gag aaa	cac aa	a gtc	tac	gcc	tgc	gaa	gtc	acc	cat	cag	1056
Ala Asp Tyr												
	340			345					350			
ggc ctg ago	tcg ccc	qtc ac	a aaq	agc	ttc	aac	agg	gga	gag	tct		1101
Gly Leu Ser	_	_	_	_								
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4222 Dogge		e		C		s. 11 a	- L DM-	MOT	7 A II	- 4		i single shein
	_	f Artif	icial	Seq	ience	e: "s	shPM	L-MCI	H4",	a de	esigne	d single chain
<223> Descr Fv gene seq	_	f Artif	icial	Seqi	ience	e: ":	shPMI	L-MCI	H4",	a de	esigned	d single chain
Fv gene seq	_	f Artif	icial	Seq	ience	e: ":	shPMI	L-MCI	H4",	a de	esigne	d single chain
Fv gene seq	uence											d single chain
Fv gene seq	uence agc tgt	atc at	c ctc	ttc	ttg	gta	gca	aca	gct	aca	ggt	
Fv gene seq <400> 27 atg gga tgg	uence agc tgt	atc at	c ctc	ttc	ttg	gta	gca	aca	gct	aca	ggt	48
Fv gene seq <400> 27 atg gga tgg Met Gly Trp 1	uence agc tgt Ser Cys 5	atc at Ile Il	c ctc e Leu	ttc Phe	ttg Leu 10	gta Val	gca Ala	aca Thr	gct Ala	aca Thr 15	ggt Gly	48
Fv gene seq <400> 27 atg gga tgg Met Gly Trp	agc tgt Ser Cys 5	atc at Ile Il	c ctc e Leu g cag	ttc Phe	ttg Leu 10	gta Val ggt	gca Ala cca	aca Thr	gct Ala ctt	aca Thr 15	ggt Gly aga	48
<pre>Fv gene seq &lt;400&gt; 27 atg gga tgg Met Gly Trp 1 gtc gac tcc</pre>	agc tgt Ser Cys 5	atc at Ile Il	c ctc e Leu g cag	ttc Phe	ttg Leu 10	gta Val ggt	gca Ala cca	aca Thr	gct Ala ctt	aca Thr 15	ggt Gly aga	48
<pre>Fv gene seq &lt;400&gt; 27 atg gga tgg Met Gly Trp 1 gtc gac tcc Val Asp Ser</pre>	agc tgt Ser Cys 5 cag gtc Gln Val	atc at Ile Il caa ct Gln Le	c ctc e Leu g cag u Gln	ttc Phe gag Glu 25	ttg Leu 10 agc Ser	gta Val ggt Gly	gca Ala cca Pro	aca Thr ggt Gly	gct Ala ctt Leu 30	aca Thr 15 gtg Val	ggt Gly aga Arg	48 96
<pre>Fv gene seq &lt;400&gt; 27 atg gga tgg Met Gly Trp 1 gtc gac tcc</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg	atc at Ile Il caa ct Gln Le	c ctc e Leu g cag u Gln	ttc Phe gag Glu 25	ttg Leu 10 agc Ser	gta Val ggt Gly	gca Ala cca Pro	aca Thr ggt Gly	gct Ala ctt Leu 30	aca Thr 15 gtg Val	ggt Gly aga Arg	48
<pre>Fv gene seq &lt;400&gt; 27 atg gga tgg Met Gly Trp 1 gtc gac tcc Val Asp Ser cct agc cag</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg	atc at Ile Il caa ct Gln Le	c ctc e Leu g cag u Gln	ttc Phe gag Glu 25	ttg Leu 10 agc Ser	gta Val ggt Gly	gca Ala cca Pro	aca Thr ggt Gly	gct Ala ctt Leu 30	aca Thr 15 gtg Val	ggt Gly aga Arg	48 96
<pre>Fv gene seq  &lt;400&gt; 27 atg gga tgg Met Gly Trp     1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln     35</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg	atc at Ile Il caa ct Gln Le agc ct Ser Le	c ctc e Leu g cag u Gln g acc u Thr 40	ttc Phe gag Glu 25 tgc Cys	ttg Leu 10 agc Ser acc	gta Val ggt Gly gtg Val	gca Ala cca Pro tct Ser	aca Thr ggt Gly ggc Gly 45	gct Ala ctt Leu 30 tac Tyr	aca Thr 15 gtg Val tca Ser	ggt Gly aga Arg att	48 96 144
<pre>Fv gene seq  &lt;400&gt; 27 atg gga tgg Met Gly Trp     1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg Thr Leu cat gcc	atc at Ile Il caa ct Gln Le agc ct Ser Le	c ctc e Leu g cag u Gln g acc u Thr 40	ttc Phe gag Glu 25 tgc Cys	ttg Leu 10 agc Ser acc Thr	gta Val ggt Gly gtg Val	gca Ala cca Pro tct Ser	aca Thr ggt Gly ggc Gly 45	gct Ala ctt Leu 30 tac Tyr	aca Thr 15 gtg Val tca ser	ggt Gly aga Arg att Ile	48 96
<pre>Fv gene seq  &lt;400&gt; 27 atg gga tgg Met Gly Trp     1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln     35 acc agc gat</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg Thr Leu cat gcc	atc at Ile Il caa ct Gln Le agc ct Ser Le	c ctc e Leu g cag u Gln g acc u Thr 40 c tgg	ttc Phe gag Glu 25 tgc Cys	ttg Leu 10 agc Ser acc Thr	gta Val ggt Gly gtg Val	gca Ala cca Pro tct Ser	aca Thr ggt Gly ggc Gly 45	gct Ala ctt Leu 30 tac Tyr	aca Thr 15 gtg Val tca ser	ggt Gly aga Arg att Ile	48 96 144
<pre> &lt;400&gt; 27 atg gga tgg Met Gly Trp     1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln     35  acc agc gat Thr Ser Asp     50</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg Thr Leu cat gcc His Ala	atc at Ile Il caa ct Gln Le agc ct Ser Le tgg ag Trp Se	c ctc e Leu g cag u Gln g acc u Thr 40 c tgg r Trp	ttc phe gag Glu 25 tgc Cys	ttg Leu 10 agc Ser acc Thr	gta Val ggt Gly gtg Val cag	gca Ala cca Pro tct Ser cca Pro 60	aca Thr ggt Gly ggc Gly 45 cct Pro	gct Ala ctt Leu 30 tac Tyr	aca Thr 15 gtg Val tca Ser cga Arg	ggt Gly aga Arg att Ile	48 96 144 192
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<pre> &lt;400&gt; 27 atg gga tgg Met Gly Trp     1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln     35  acc agc gat Thr Ser Asp     50</pre>	agc tgt Ser Cys 5 cag gtc Gln Val 20 acc ctg Thr Leu cat gcc His Ala	atc at Ile Il caa ct Gln Le agc ct Ser Le tgg ag Trp Se 5	c ctc e Leu g cag u Gln g acc u Thr 40 c tgg r Trp t	ttc Phe gag Glu 25 tgc Cys gtt Val	ttg Leu 10 agc Ser acc Thr cgc Arg	gta Val ggt Gly gtg Val cag Gln	gca Ala cca Pro tct Ser cca Pro 60	aca Thr ggt Gly ggc Gly 45 cct Pro	gct Ala ctt Leu 30 tac Tyr gga Gly	aca Thr 15 gtg Val tca ser cga Arg	ggt Gly aga Arg att Ile ggt Gly	48 96 144 192
<pre>Fv gene seq  &lt;400&gt; 27 atg gga tgg Met Gly Trp 1  gtc gac tcc Val Asp Ser  cct agc cag Pro Ser Gln</pre>	agc tgt Ser Cys Scag gtc Gln Val 20 acc ctg Thr Leu cat gcc His Ala att gga Ile Gly	atc at Ile Il caa ct Gln Le agc ct Ser Le tgg ag Trp Se 5 tac at Tyr Il 70	c ctc e Leu g cag u Gln g acc u Thr 40 c tgg r Trp t agt	ttc Phe gag Glu 25 tgc Cys gtt Val tat Tyr	ttg Leu 10 agc Ser acc Thr cgc Arg	gta Val ggt Gly gtg Val cag Gln gga Gly 75	gca Ala cca Pro tct Ser cca Pro 60 atc	aca Thr ggt Gly ggc Gly 45 cct Pro	gct Ala ctt Leu 30 tac Tyr gga Gly acc Thr	aca Thr 15 gtg Val tca Ser cga Arg tat Tyr	ggt Gly aga Arg att Ile ggt Gly aat Asn 80	48 96 144 192 240
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85 90 95

_		_	_	_		-	-			_	_	gac Asp				336
												atg Met 125				384
												ggt Gly				432
												atg Met				480
			_		_					_		acc Thr				528
_	-	-	_	-		-	-		_		-	tac Tyr				576
												tcc Ser 205				624
	~~			_	~		_		_		_	ggt Gly		_		672
					_					_		gct Ala				720
												caa Gln				768
	_				_	_				_	_	tac Tyr	_	_		816
												acc Thr 285				864
_		-						_	_			cag Gln		_	_	912
												agc Ser				960

```
cet gag eec cag gee eea gge egg tac tte gee cae age ate etg ace
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Pro Glu Pro Gln Ala Pro Gly Arg Tyr Phe Ala His Ser Ile Leu Thr
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                325
                                                                   1056
gtg tcc gaa gag gaa tgg aac acg ggg gag acc tac acc tgc gtg gcc
Val Ser Glu Glu Glu Trp Asn Thr Gly Glu Thr Tyr Thr Cys Val Ala
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                                345
                                                                   1104
cat gag gcc ctg ccc aac agg gtc acc gag agg acc gtg gac aag tcc
His Glu Ala Leu Pro Asn Arg Val Thr Glu Arg Thr Val Asp Lys Ser
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                            360
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Thr Glu Gly Glu Val Ser Ala Asp Glu Glu Gly Phe Glu
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                        375
<210> 28
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<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: "EF-1", an artificially synthesized
primer sequence
<400> 28
                                                                   18
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<210> 29
<211> 107
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: "SCP-C2", an artificially synthesized
primer sequence
<400> 29
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<212> DNA
<213> Artificial Sequence
<220>
<221> CDS
<222> (1)..(1557)
<220>
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<223> Description of Artificial Sequence: "shPM1( $\Delta$ EL)-BvGS3", a designed single chain Fv gene sequence

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				Cys												10
1	OLY	ııp	Jer	5	110		Бец	E 110	10	vuz	2324			15	OL)	
				,					10					10		
atc	gac	tcc	cag	gtc	caa	cta	caq	gag	age	aat	cca	aat.	ctt	ata	aga	96
				Val												
			20				02	25		1		1	30		3	
cct	адс	caq	acc	ctg	ачс	cta	acc	tac	acc	ata	tct	qqc	tac	tca	att	144
				Leu												
		35					40	•				45				
acc	agc	gat	cat	gcc	tgg	agc	tgg	gtt	cgc	cag	cca	cct	gga	cga	ggt	192
Thr	Ser	Asp	His	Ala	Trp	Ser	Trp	Val	Arg	Gln	Pro	Pro	Gly	Arg	Gly	
	50					55					60					
ctt	gag	tgg	att	gga	tac	att	agt	tat	agt	gga	atc	aca	acc	tat	aat	240
Leu	Glu	Trp	Ile	Gly	Tyr	Ile	Ser	Tyr	Ser	Gly	Ile	Thr	Thr	Tyr	Asn	
65					70					75					80	
				tcc	-			_	_	_	_		_			288
Pro	Ser	Leu	Lys	Ser	Arg	Val	Thr	Met		Arg	Asp	Thr	Ser		Asn	
				85					90					95		
																226
				aga												336
GIN	Pne	ser		Arg	ьeu	ser	ser		Thr	Ala	Ala	Asp		Ara	vai	
			100					105					110			
tat	+ > +	+~+	aa.	aga	taa	c+ a	act	caa	act	3.CG	act	ato	aac	tac	taa	384
				Arg												301
TYL	ı yı	115	лта	Arg	267	пец	120	nr9	1111	1111	AΙα	125	TID D	-7-		
							120					100				
aat.	caa	aac	agg	ctc	atic	aca	ata	tcc	t.ca	aat.	aat	aat	aat	t.ca	aat	432
				Leu												
1	130	1				135				2	140	2	2		•	
											_					
ggt	ggt	ggt	tcg	ggt	ggt	ggc	gga	tcg	gac	atc	cag	atg	acc	cag	agc	480
				Gly												
145	•	•		-	150	-	-		_	155					160	
cca	agc	agc	ctg	agc	gcc	agc	gtg	ggt	gac	aga	gtg	acc	atc	acc	tgt	528
Pro	Ser	Ser	Leu	Ser	Ala	Ser	Val	Gly	Asp	Arg	Val	Thr	Ile	Thr	Cys	
				165		•			170					175		
				gac												576
Arg	Ala	Ser	Gln	Asp	Ile	Ser	Ser	Tyr	Leu	Asn	Trp	Tyr	Gln	Gln	Lys	
			180					185					190			
				cca												624
Pro	Gly	-	Ala	Pro	Lys	Leu		Ile	Tyr	Tyr	Thr		Arg	Leu	His	
		195					200					205				

	t ggt r Gly 210															672
	c ttc r Phe 5															720
	c caa s Gln															768
	g gaa 1 Glu															816
	c gga y Gly															864
	t gtg u Val 290															912
	c tca r Ser 5			_	_		_		_			-	_			960
	a cga y Arg															1008
	c tat r Tyr															1056
	c aag r Lys															1104
	c gcg r Ala 370															1152
	c tac p Tyr 5															1200
	t tcg y Ser															1248
	c cag r Gln															1296
at	c acc	tgt	aga	gcc	agc	cag	gac	atc	agc	agt	tac	ctg	aat	tgg	tac	1344

Ile Thr Cys Arg Ala Ser Gln Asp Ile Ser Ser Tyr Leu Asn Trp Tyr 435 440 cag cag aag cca gga aag gct cca aag ctg ctg atc tac tac acc tcc 1392 Gln Gln Lys Pro Gly Lys Ala Pro Lys Leu Leu Ile Tyr Tyr Thr Ser aga ctg cac tct ggt gtg cca agc aga ttc agc ggt agc ggt agc ggt 1440 Arg Leu His Ser Gly Val Pro Ser Arg Phe Ser Gly Ser Gly Ser Gly 465 470 475 ace gac ttc acc ttc acc atc age age ctc cag cca gag gac atc gct Thr Asp Phe Thr Phe Thr Ile Ser Ser Leu Gln Pro Glu Asp Ile Ala 485 490 495 acc tac tac tgc caa cag gga aat act tta cca tac acg ttc ggc caa Thr Tyr Tyr Cys Gln Gln Gly Asn Thr Leu Pro Tyr Thr Phe Gly Gln 500 505 1557 ggg acc aag gtg gaa atc aaa Gly Thr Lys Val Glu Ile Lys 515 <210> 31 <211> 29 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: "Kappal", an artificially synthesized primer sequence <400> 31 29 ccgccatctg atgagcagtt gaaatctgg <210> 32 <211> 54 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: "Kappa2", an artificially synthesized primer sequence <400> 32 ttatttatcg tcatcgtctt tgtagtcaag cttagactct cccctgttga agct 54 <210> 33 <211> 29 <212> DNA <213> Artificial Sequence <220>

<223> Description of Artificial Sequence: "SCP-K", an artificially synthesized

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primer sequence
<400> 33
                                                                   29
ttcaactgct catcagatgg cgggaagat
<210> 34
<211> 1878
<212> DNA
<213> Artificial Sequence
<220>
<221> CDS
<222> (1)..(1878)
<220>
<223> Description of Artificial Sequence: "shPM1-Kappa-BvGS3", a designed single
chain Fv gene sequence
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Met Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly
                                     10
gtc gac tcc cag gtc caa ctg cag gag agc ggt cca ggt ctt gtg aga
                                                                   96
Val Asp Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Arg
             20
                                                                   144
cet age cag ace etg age etg ace tge ace gtg tet gge tae tea att
Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Tyr Ser Ile
         35
acc agc gat cat gcc tgg agc tgg gtt cgc cag cca cct gga cga ggt
                                                                   192
Thr Ser Asp His Ala Trp Ser Trp Val Arg Gln Pro Pro Gly Arg Gly
     50
ctt gag tgg att gga tac att agt tat agt gga atc aca acc tat aat
                                                                   240
Leu Glu Trp Ile Gly Tyr Ile Ser Tyr Ser Gly Ile Thr Thr Tyr Asn
 65
                     70
                                                                   288
cca tct ctc aaa tcc aga gtg aca atg ctg aga gac acc agc aag aac
Pro Ser Leu Lys Ser Arg Val Thr Met Leu Arg Asp Thr Ser Lys Asn
cag ttc agc ctg aga ctc agc agc gtg aca gcc gcc gac acc gcg gtt
                                                                   336
Gln Phe Ser Leu Arg Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val
            100
                                105
                                                                   384
tat tat tgt gca aga tcc cta gct cgg act acg gct atg gac tac tgg
Tyr Tyr Cys Ala Arg Ser Leu Ala Arg Thr Thr Ala Met Asp Tyr Trp
        115
                            120
ggt caa ggc agc ctc gtc aca gtc tcc tca ggt ggt ggt tcg ggt
                                                                   432
Gly Gln Gly Ser Leu Val Thr Val Ser Ser Gly Gly Gly Ser Gly
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140

130

135

ggt Gly 145	ggt Gly	ggt Gly	tcg Ser	ggt Gly	ggt Gly 150	ggc Gly	gga Gly	tcg Ser	gac Asp	atc Ile 155	cag Gln	atg Met	acc Thr	cag Gln	agc Ser 160	480
				agc Ser 165											Cys	528
				gac Asp												576
		_	_	cca Pro	_	_	_						_			624
				agc Ser												672
				agc Ser	_		_		_	_		_				720
				aac Asn 245												768
	_			ggt Gly				_					_			816
				gac Asp												864
		_		agc Ser	_		_	_	_		_					912
				agc Ser												960
				gag Glu 325												1008
				tct Ser												1056
				ttc Phe												1104

acc Thr	gcg Ala 370	gtt Val	tat Tyr	tat Tyr	tgt Cys	gca Ala 375	aga Arg	tcc Ser	cta Leu	gct Ala	cgg Arg 380	act Thr	acg Thr	gct Ala	atg Met	1152
gac Asp 385	tac Tyr	tgg Trp	ggt Gly	caa Gln	ggc Gly 390	agc Ser	ctc Leu	gtc Val	aca Thr	gtc Val 395	tcc Ser	tca Ser	ggt Gly	ggt Gly	ggt Gly 400	1200
					ggt Gly											1248
					agc Ser											1296
					agc Ser											1344
					aag Lys											1392
					gtg Val 470											1440
	_				acc Thr		-	_								1488
					cag Gln											1536
					atc Ile											1584
					gat Asp											1632
					aac Asn 550											1680
					ctc Leu											1728
gag Glu	cag Gln	gac Asp	agc Ser 580	aag Lys	gac Asp	agc Ser	acc Thr	tac Tyr 585	agc Ser	ctc Leu	agc Ser	agc Ser	acc Thr 590	ctg Leu	acg Thr	1776
ctg	agc	aaa	gca	gac	tac	gag	aaa	cac	aaa	gtc	tac	gcc	tgc	gaa	gtc	1824

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Leu Ser Lys Ala Asp Tyr Glu Lys His Lys Val Tyr Ala Cys Glu Val
                            600
        595
acc cat cag ggc ctg agc tcg ccc gtc aca aag agc ttc aac agg gga
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Thr His Gln Gly Leu Ser Ser Pro Val Thr Lys Ser Phe Asn Arg Gly
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                                                                   1878
gag tct
Glu Ser
625
<210> 35
<211> 29
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: "MCH4-1", an artificially synthesized
primer sequence
<400> 35
                                                                   29
gtggaaatca aagtggccct gcacaggcc
<210> 36
<211> 68
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: "MCH4-2.1", an artificially
synthesized primer sequence
<400> 36
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                                                                    68
acggtcct
<210> 37
<211> 29
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: "SCP-Mu", an artificially synthesized
primer sequence
<400> 37
                                                                    29
tgcagggcca ctttgatttc caccttggt
<210> 38
<211> 53
<212> DNA
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<213> Artificial Sequence

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<223> Description of Artificial Sequence: "MCH4-2.2", an artificially
synthesized primer sequence
aaagcggccg cttattattt atcgtcatcg tctttgtagt caagcttctc aaa
<210> 39
<211> 1920
<212> DNA
<213> Artificial Sequence
<220>
<221> CDS
<222> (1)..(1920)
<220>
<223> Description of Artificial Sequence: "shPM1-MCH4-BvGS3", a designed single
chain Fv gene sequence
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Met Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr Gly
gtc gac tcc cag gtc caa ctg cag gag agc ggt cca ggt ctt gtg aga
                                                                  96
Val Asp Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly Leu Val Arg
             20
cct age cag ace etg age etg ace tge ace gtg tet gge tac tea att
                                                                  144
Pro Ser Gln Thr Leu Ser Leu Thr Cys Thr Val Ser Gly Tyr Ser Ile
         35
acc age gat cat gee tgg age tgg gtt ege cag cca eet gga ega ggt
                                                                  192
Thr Ser Asp His Ala Trp Ser Trp Val Arg Gln Pro Pro Gly Arg Gly
     50
ctt gag tgg att gga tac att agt tat agt gga atc aca acc tat aat
Leu Glu Trp Ile Gly Tyr Ile Ser Tyr Ser Gly Ile Thr Thr Tyr Asn
65
cca tct ctc aaa tcc aga gtg aca atg ctg aga gac acc agc aag aac
                                                                  288
Pro Ser Leu Lys Ser Arg Val Thr Met Leu Arg Asp Thr Ser Lys Asn
                 85
cag ttc agc ctg aga ctc agc agc gtg aca gcc gcc gac acc gcg gtt
                                                                  336
Gln Phe Ser Leu Arg Leu Ser Ser Val Thr Ala Ala Asp Thr Ala Val
            100
                                105
tat tat tgt gca aga tcc cta gct cgg act acg gct atg gac tac tgg
                                                                  384
Tyr Tyr Cys Ala Arg Ser Leu Ala Arg Thr Thr Ala Met Asp Tyr Trp
        115
                            120
ggt caa ggc agc ctc gtc aca gtc tcc tca ggt ggt ggt tcg ggt
                                                                  432
Gly Gln Gly Ser Leu Val Thr Val Ser Ser Gly Gly Gly Ser Gly
    130
                        135
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ggt ggt tcg ggt ggt ggc gga tcg gac atc cag atg acc cag agc 480  Gly Gly Gly Ser Gly Gly Gly Ser Asp Ile Gln Met Thr Gln Ser 160  150
145  cca agc agc ctg agc gcc agc gtg ggt gac aga gtg acc atc acc tgt 528  cca agc agc ctg agc gcc agc gtg ggt gac aga gtg acc atc acc tgt 528  cca agc agc ctg agc gcc agc gtg ggt gac aga gtg acc atc acc tgt 528  cca agc agc ctg agc gcc agc gtg ggt gac aga gtg acc atc acc tgt 528  165  170  175
pro Ser Ser Head 505 170  165  165  170  170  170  170  170  170  170  17
Arg Ala Sei Gin 135 185
cca gga aag gct cca aag ctg ctg atc tac tac acc tcc aga ctg cdo  cca gga aag gct cca aag ctg ctg atc tac tac acc tcc aga ctg cdo  Pro Gly Lys Ala Pro Lys Leu Leu Ile Tyr Tyr Thr Ser Arg Leu His  205
tot ggt gtg cca agc aga ttc agc ggt agc ggt acc gac ttc 672  tct ggt gtg cca agc aga ttc agc ggt agc ggt acc gac ttc 672  tct ggt gtg cca agc aga ttc agc ggt agc ggt acc gac ttc 672  Ser Gly Val Pro Ser Arg Phe Ser Gly Ser Gly Thr Asp Phe 220
acc ttc acc atc agc agc ctc cag cca gag gac atc gct acc tac tac 720  acc ttc acc atc agc agc ctc cag cca gag gac atc gct acc tac tac 720  Thr Phe Thr Ile Ser Ser Leu Gln Pro Glu Asp Ile Ala Thr Tyr Tyr  240  230
230 225 230 230 235 235 236 237 268
Cys Gln Gin Giy And 250
gtg gaa atc aaa ggt ggt ggt tcg ggt ggt ggt ggt tcg ggt ggt
ggc gga tcg gtc gac tcc cag gtc caa ctg cag gag agc ggt cca ggt solv Gly Gly Ser Val Asp Ser Gln Val Gln Leu Gln Glu Ser Gly Pro Gly 280
ctt gtg aga cct agc cag acc ctg agc ctg acc tgc acc gtg tct ggc 912  ctt gtg aga cct agc cag acc ctg agc ctg acc tgc acc gtg tct ggc 912  ctt gtg aga cct agc cag acc ctg agc ctg acc tgc acc gtg tct ggc 912
tac tca att acc agc gat cat gcc tgg agc tgg gtt cgc cag cca cct  tac tca att acc agc gat cat gcc tgg agc tgg gtt cgc cag cca cct  Tyr Ser Ile Thr Ser Asp His Ala Trp Ser Trp Val Arg Gln Pro  320  310
305 agt gga atc aca 1008
Gly Arg Gly Bed 325 330
acc tat aat cca tct ctc aaa tcc aga gtg aca atg ctg aga gac acc 1056  Thr Tyr Asn Pro Ser Leu Lys Ser Arg Val Thr Met Leu Arg Asp Thr  350  340  345
agc aag aac cag ttc agc ctg aga ctc agc agc gtg aca gcc gcc gac 1104  Ser Lys Asn Gln Phe Ser Leu Arg Leu Ser Ser Val Thr Ala Ala Asp  365  360

acc Thr	gcg Ala 370	gtt Val	tat Tyr	tat Tyr	tgt Cys	gca Ala 375	aga Arg	tcc Ser	cta Leu	gct Ala	cgg Arg 380	act Thr	acg Thr	gct Ala	atg Met	1152
gac Asp 385	tac Tyr	tgg Trp	ggt Gly	caa Gln	ggc Gly 390	agc Ser	ctc Leu	gtc Val	aca Thr	gtc Val 395	tcc Ser	tca Ser	ggt Gly	ggt Gly	ggt Gly 400	1200
								ggt Gly								1248
								gcc Ala 425								1296
								atc Ile								1344
cag Gln	cag Gln 450	aag Lys	cca Pro	gga Gly	aag Lys	gct Ala 455	cca Pro	aag Lys	ctg Leu	ctg Leu	atc Ile 460	tac Tyr	tac Tyr	acc Thr	tcc Ser	1392
								aga Arg								1440
								agc Ser								1488
acc Thr	tac Tyr	tac Tyr	tgc Cys 500	caa Gln	cag Gln	gga Gly	aat Asn	act Thr 505	tta Leu	cca Pro	tac Tyr	acg Thr	ttc Phe 510	ggc	caa Gln	1536
								gcc Ala								1584
								ctg Leu								1632
								tct Ser								1680
								tcc Ser								1728
								cca Pro 585								1776
atc	ctg	acc	gtg	tcc	gaa	gag	gaa	tgg	aac	acg	ggg	gag	acc	tac	acc	1824

I1	e	Leu	Thr 595	Val	Ser	Glu	Glu	Glu 600	Trp	Asn	Thr	Gly	Glu 605	Thr	Tyr	Thr	
to C)	JC /S	gtg Val 610	gcc Ala	cat His	gag Glu	gcc Ala	ctg Leu 615	ccc Pro	aac Asn	agg Arg	gtc Val	acc Thr 620	gag Glu	agg Arg	acc Thr	gtg Val	1872
A	ac sp 25	-	tcc Ser	acc Thr	gag Glu	630 ggg	014	gtg Val	agc Ser	gcc Ala	gac Asp 635	gag Glu	gag	ggc	ttt Phe	gag Glu 640	1920